

Pending Claims as of November 16, 2001

76. An HLA-DR typing process comprising the steps of:

- (a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:
- (i) DNA sequences encoding amino acids 8-14 of said locus;
  - (ii) DNA sequences encoding amino acids 26-32 of said locus;
  - (iii) DNA sequences encoding amino acids 72-78 of said locus;
  - (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
  - (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

77. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

78. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

79. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

80. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCAATTCTCAATGGGACGGAGCGGGTGCGGTTCTGGAGA  
GACACTTCATAACCAGGAGGAGTACGCGCCTCGACAGCG  
ACGTGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCAGGGCCAGGTGGACAATTACTGCAGACACAACATACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAAATGGAGACTGGACCTC  
CAGACCTGGTGTAGCTAGAAACATTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGCTTGTGCTGGGCCTGCTC  
TTCCTGGGGCCGGGCTGTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (ii) GGGGACACCCGACCACGTTCTTGGAGCAGGTTAACATGAG  
TGTCAATTCTCAACGGGACGGAGCGGGTGCGGTTCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTCGACAGC  
GACGTGGGGAGTACCGGGCCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCAGGGCCGGTGGACACCTACTGCAGACACAACATACGGG  
GTTGGTGAGAGCTTCACAGTGCAGCGGCAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCTGCAGCACAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAAATGGAGACTGGACC  
TTCCAGACCCCTGGTGTAGCTGGAAACAGTTCTCGGAGTGGGA  
GAGGTTTACACCTCCAAGTGGAGCACGGTCTGAATCTGCACAG  
CCTCTCACAGTGAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGCTTCGTGCTGGGCCTG  
CTCTCCTGGGGCCGGGCTGTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (iii) a DNA sequence which is fully complementary to the DNA sequence of (i) or (ii); and
- (iv) a DNA sequence which differs from the DNA sequence of (i) or (ii) in codon sequence due to the degeneracy of the genetic code, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

81. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTCTTGAGCTGCTTAAGTCTGAG  
TGTCAATTCTCAATGGACGGAGCGGGTGCCTGAGA  
GACACTTCATAACCAGGAGGAGTACGCGCCTCGACAGCG  
ACGTGGGGAGTACCGGGCGGTGAGGGAGCTGGGCGGCCTG  
ATGCCAGACTGGAACAGCAGAAGGACCTCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACACTACGGGG  
TTGTGGAGAGCTTCACAGTGCAGCGGCAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCCTGCAGCACCAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTCCGGAACGGCAGGAAGAGAAGGGCTGGG  
GTGGTGTCCACGGGCGGTGATCCAGAAATGGAGACTGGACCTTC  
CAGACCCCTGGTGTGCTAGAAACATTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGCTTGTGCTGGGCGTGTGTC  
TTCCTGGGCGGGCTGTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (ii) GGGGACACCCGACCACGTTCTTGAGCAGGTTAAACATGAG  
TGTCAATTCTCAACGGACGGAGCGGGTGCCTGAG  
AGATACTTCTATCACCAAGAGGGAGTACGTGCCTCGACAGC  
GACGTGGGGAGTACCGGGCGGTGACGGAGCTGGGCGGCCT  
GATGCCAGTACTGGAACAGCCAGAAGGACCTCTGGAGCAG  
AAGCGGGCCGCGGTGGACACCTACTGCAGACACAACACTACGGG  
GTTGGTGTGAGACTTCACAGTGCAGCGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCCTGCAGCACCA  
AACCTCTGGTCTGCTCTGTGAATGGTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCGTGTGAAACAGTCTCGGAGTGGACC  
TTCCAGACCCCTGGTGTGAGTGGAAACAGTCTCGGAGTGG  
GAGGTTTACACCTCCAAGTGGAGCACGGTCTGAATCTGCACAG  
CCTCTCACAGTGGAAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGCTTCGTGCTGGGCGT  
CTCTCCTGGGGCGGGCTGTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (iii) a DNA sequence which is fully complementary to the DNA sequence of (i) or (ii); and
- (iv) a DNA sequence which differs from the DNA sequence of (i) or (ii) in codon sequence due to the degeneracy of the genetic code, and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

83. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

84. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

85. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:
- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and
- (d) detecting areas of hybridization between said DNA to be typed and said second DNA.

86. The HLA-DR typing process according to claim 76 or 78, wherein said DNA sequence is characterized by a nucleotide sequence selected from the group consisting of:

TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA,  
GGGGCCAGGTGGACAATTA, TGGAGCAGGTTAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

87. The HLA-DR typing process according to claim 77 or 79, wherein said second DNA is characterized by a nucleotide sequence selected from the group consisting of:  
TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA, GGGGCCAGGTGGACAATTA,  
TGGAGCAGGTTAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

88. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said DNA sequence.

89. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said second DNA.

90. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said DNA in said sample to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

91. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said size-fractionated DNA

to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

92. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein said DNA sequence is a labeled DNA sequence and its label is used for detecting hybridization between said DNA in said sample and said DNA sequence.

93. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein said second DNA is a labeled DNA and its label is used for detecting hybridization between said size-fractionated DNA and said second DNA.

94. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (ii) DNA sequences encoding amino acids 26-32 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iii) DNA sequences encoding amino acids 72-78 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

95. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

96. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of

hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences.

97. The HLA-DR typing kit according to any one of claims 94, 95 or 96, wherein said DNA sequence is labeled.

98. The HLA-DR typing kit according to any one of claims 94, 95 or 96, further comprising a 19-mer hybridization control, said hybridization control being a DNA being the nucleotide sequence: GCTTCGACAGCGACGTGGG.

99. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of an HLA-DR- $\beta$  locus of the human lymphocyte antigen complex, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

100. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

101. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting

- essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C, and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences.

102. The HLA-DR typing kit according to anyone of claims 99, 100 or 101, wherein said DNA sequence is labeled.

Pending Claims as of April 2002

76. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

77. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;  
(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

78. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

79. An HLA-DR typing process comprising the steps of:

(a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

(b) size-fractionating said restricted DNA;  
(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex to allow determination of one or more HLA-DR alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

80. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being selected from the group consisting of:

- (i) GGGGACACCGACCACGTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCAATTCTCAATGGGACGGAGCGGGTGCCTGGAGA  
GACACTTCCATAACCAGGAGGAGTACGCGCCTCGACAGCG  
ACGTGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCAGACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCGGGGCCAGGTGGACAATTACTGCAGACACAACATACGGGG  
TGAGTGGAGAGCTTCACAGTGCAGCGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTTCCGGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAAATGGAGACTGGACCTC  
CAGACCCCTGGTGTAGCTAGAAACATTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAAATGGAGTGCACGGCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGCTTGTGCTGGGCCTGCTC  
TTCCTGGGGCCGGCTGTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (ii) GGGGACACCGACCACGTTCTTGGAGCAGGTTAACATGAG  
TGTCAATTCTCAACGGGACGGAGCGGGTGCCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGAGTACCGGGCGGTGACGGAGCTGGGGCGGCCT  
GATGCCAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGGTGGACACCTACTGCAGACACAACATACGGG  
GTTGGTGAGAGCTTCACAGTGCAGCGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCTGCAGCACAC  
AACCTCCTGGTCTGCTGTGAATGGTTCTATCCAGGCAGC  
ATTGAAGTCAGGTGGTTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAAATGGAGACTGGACC  
TTCCAGACCCCTGGTGTAGCTGGAAACAGTTCCTCGGAGTGG  
GAGGTTTACACCTCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGGAAATGGAGAGCAGGCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGCTTCGTGCTGGGCCTG  
CTCTCCTGGGGCCGGCTGTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (iii) a DNA sequence which is fully complementary to the DNA sequence of (i) or (ii); and
- (iv) a DNA sequence which differs from the DNA sequence of (i) or (ii) in codon sequence due to the degeneracy of the genetic code, and

(b) detecting areas of hybridization between said DNA in said sample and said DNA sequence.

81. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTCTTGGAGCTGCTTAAGTCTGAG  
TGTCAATTCTCAATGGGACGGAGCGGGTGCGGTTCTGGAGA  
GACACTTCATAACCAGGAGGAGTACGCGCCTCGACAGCG  
ACGTGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAGA  
AGCAGGGCCAGGTGGACAATTACTGCAGACACAACATACGGGG  
TTGTGGAGAGCTCACAGTGCAGCGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTCCCGAACGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTC  
CAGACCCCTGGTGTAGCTAGAACATTTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAATGGAGTGCACGGCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGCTTGTGCTGGGCCTGCTC  
TTCCTGGGGCCGGCTGTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (ii) GGGGACACCCGACCACGTTCTTGGAGCAGTTAACATGAG  
TGTCAATTCTCAACGGGACGGAGCGGGTGCGGTTCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGAGTACCGGGCGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGGTGGACACCTACTGCAGACACAACATACGGG  
GTTGGTGGAGAGCTTCACAGTGCAGCGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTCTATCCAGGCAGC  
ATTGAAGTCAGTGGTCCCGAACGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCCTGGTGTAGCTGGAAACAGTCCCTCGGAGTGG  
GAGGTTTACACCTCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGGAATGGAGAGCACGGCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGCTTCGTGCTGGCCTG  
CTCTCCTGGGGCCGGCTGTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (iii) a DNA sequence which is fully complementary to the DNA sequence of (i) or (ii); and
- (iv) a DNA sequence which differs from the DNA sequence of (i) or (ii) in codon sequence due to the degeneracy of the genetic code, and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being selected from the group consisting of:

- (i) GGGGACACCCGACCACGTTCTTGAGCTGCTTAAGTCTGAG  
TGTCAATTCTCAATGGGACGGAGCGGGTGCCTGGAGA  
GACACTTCATAACCAGGAGGAGTACCGCGCTTCGACAGCG  
ACGTGGGGAGTACCGGGCGGTGAGGGAGCTGGGGCGGCCTG  
ATGCCGAGTACTGGAACAGCAGAAGGACCTCCTGGAGCAGA  
AGCAGGGCCAGGTGGACAATTACTGCAGACACAACATACGGGG  
TTGTGGAGAGCTCACAGTGCAGCGCGAGTCCATCCTCAGG  
TGACTGTGTATCCTGCAAGACCCAGCCCCTGCAGCACCACAA  
CCTCCTGGTCTGCTCTGTGAGTGGTTCTATCCAGGCAGCAT  
TGAAGTCAGTGGTCCCGAACGGCCAGGAAGAGAAGGCTGGG  
GTGGTGTCCACGGGCCTGATCCAGAATGGAGACTGGACCTTC  
CAGACCCCTGGTGTAGCTAGAAACATTCTCGGAGTGGAGAG  
GTTTACACCTGCCAAGTGGAGCACCCAAGCGTAACGAGCCCT  
CTCACAGTGGAAATGGAGTGCACGGTCTGAATCTGCACAGAGC  
AAGATGCTGAGTGGAGTCGGGGCTTGTGCTGGGCCTGCTC  
TTCCTGGGCGGGCTGTTCATCTACTTCAGGAATCAGAAA  
GGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (ii) GGGGACACCCGACCACGTTCTTGAGCAGGTTAACATGAG  
TGTCAATTCTCAACGGGACGGAGCGGGTGCCTGGAC  
AGATACTTCTATCACCAAGAGGAGTACGTGCGCTTCGACAGC  
GACGTGGGGAGTACCGGGCGGTGACGGAGCTGGGGCGGCCT  
GATGCCGAGTACTGGAACAGCCAGAAGGACCTCCTGGAGCAG  
AAGCGGGCCGGTGGACACCTACTGCAGACACAACATACGGG  
GTTGGTGGAGAGCTTCACAGTGCAGCGCGAGTCTATCCTGAG  
GTGACTGTGTATCCTGCAAAGACCCAGCCCCTGCAGCACCAC  
AACCTCCTGGTCTGCTCTGTGAATGGTTCTATCCAGGCAGC  
ATTGAAGTCAGTGGTCCGGAACGGCCAGGAAGAGAAGACT  
GGGGTGGTGTCCACAGGCCTGATCCAGAATGGAGACTGGACC  
TTCCAGACCCCTGGTGTAGCTGGAAACAGTTCCTCGGAGTGG  
GAGGTTTACACCTCCAAGTGGAGCACCCAAGCCTGACGAGC  
CCTCTCACAGTGGAAATGGAGAGCACGGTCTGAATCTGCACAG  
AGCAAGATGCTGAGTGGAGTCGGGGCTTCGTGCTGGCCTG  
CTCTCCTGGGGCGGGCTGTTCATCTACTTCAGGAATCAG  
AAAGGACACTCTGGACTTCAGCCAACAGGATTCTGAGC;
- (iii) a DNA sequence which is fully complementary to the DNA sequence of (i) or (ii); and
- (iv) a DNA sequence which differs from the DNA sequence of (i) or (ii) in codon sequence due to the degeneracy of the genetic code, and

(d) detecting hybridization between said size-fractionated DNA and said second DNA.

82. An HLA-DR typing process comprising the steps of:

(a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human

lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and
- (b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

83. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;
- (b) size-fractionating said restricted DNA;
- (c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus; and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences, and

(d) detecting areas of hybridization between said size-fractionated DNA and said second DNA.

84. An HLA-DR typing process comprising the steps of:

- (a) hybridizing DNA in a sample to be typed to a DNA sequence, said DNA sequence being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
- (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and

(b) detecting areas of hybridization between said DNA in the sample and said DNA sequence.

85. An HLA-DR typing process comprising the steps of:

- (a) restricting a first DNA isolated from an individual to be typed with at least one restriction endonuclease;

- (b) size-fractionating said restricted DNA;  
(c) hybridizing said size-fractionated DNA to be typed to a second DNA, said second DNA being capable of hybridizing to a constant region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said constant region being encoded by a DNA sequence selected from the group consisting of:
- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C; and
  - (ii) DNA sequences which are fully complementary to any of the foregoing sequences, and
- (d) detecting areas of hybridization between said DNA to be typed and said second DNA.

86. The HLA-DR typing process according to claim 76 or 78, wherein said DNA sequence is characterized by a nucleotide sequence selected from the group consisting of:

TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA,  
GGGGCCAGGTGGACAATT, TGGAGCAGGTTAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

87. The HLA-DR typing process according to claim 77 or 79, wherein said second DNA is characterized by a nucleotide sequence selected from the group consisting of:  
TGGAGCTGCTTAAGTCTGA, TCCTGGAGAGACACTTCCA, GGGGCCAGGTGGACAATT,  
TGGAGCAGGTTAACATGA, TCCTGGACAGATACTTCTA  
and GGGCCGCGGTGGACACCTA.

88. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said DNA sequence.

89. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, further comprising the step of comparing said hybridization to hybridization between DNA of known HLA-DR type and said second DNA.

90. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said DNA in said sample to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

91. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein prior to the step of detecting said areas of hybridization, the process further comprises the step of hybridizing said size-fractionated DNA

to a hybridization control, said hybridization control being a DNA having the nucleotide sequence: GCTTCGACAGCGACGTGGG.

92. The HLA-DR typing process according to any one of claims 76, 78, 80, 82 or 84, wherein said DNA sequence is a labeled DNA sequence and its label is used for detecting hybridization between said DNA in said sample and said DNA sequence.

93. The HLA-DR typing process according to any one of claims 77, 79, 81, 83 or 85, wherein said second DNA is a labeled DNA and its label is used for detecting hybridization between said size-fractionated DNA and said second DNA.

94. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (ii) DNA sequences encoding amino acids 26-32 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iii) DNA sequences encoding amino acids 72-78 of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

95. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 8-14 of said locus;
- (ii) DNA sequences encoding amino acids 26-32 of said locus;
- (iii) DNA sequences encoding amino acids 72-78 of said locus;
- (iv) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (v) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

96. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of

hybridizing to a polymorphic region of said locus to allow determination of one or more HLA alleles for use in HLA-DR- $\beta$  typing, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting essentially of amino acids 8-14, 26-32 or 72-78 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;
- (ii) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (iii) DNA sequences which are fully complementary to any of the foregoing sequences.

97. The HLA-DR typing kit according to any one of claims 94, 95 or 96, wherein said DNA sequence is labeled.

98. The HLA-DR typing kit according to any one of claims 94, 95 or 96, further comprising a 19-mer hybridization control, said hybridization control being a DNA being the nucleotide sequence: GCTTCGACAGCGACGTGGG.

99. An HLA-DR typing kit comprising a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of an HLA-DR- $\beta$  locus of the human lymphocyte antigen complex, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

100. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding amino acids 39-45 of said locus, and
- (ii) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

101. An HLA-DR typing kit comprising a DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing to a conserved region of said locus to allow determination of a HLA-DR- $\beta$  chain for use in HLA-DR- $\beta$  typing, said conserved region being encoded by a DNA sequence selected from the group consisting of:

- (i) DNA sequences encoding a majority of the amino acid sequence in a region consisting

essentially of amino acids 39-45 of a polypeptide sequence coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C, and  
(ii) DNA sequences which are fully complementary to any of the foregoing sequences.

102. The HLA-DR typing kit according to anyone of claims 99, 100 or 101, wherein said DNA sequence is labeled.

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27. An isolated DNA sequence which hybridizes to an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being capable of hybridizing, at to a polymorphic region of said locus to allow determination of one or more HLA alleles, said polymorphic region being encoded by a DNA sequence selected from the group consisting of:

- (a) DNA sequences encoding amino acids 8-14 of said locus;
- (b) DNA sequences encoding amino acids 26-32 of said locus;
- (c) DNA sequences encoding amino acids 72-78 of said locus;
- (d) DNA sequences which are portions of any one of the foregoing DNA sequences and which are capable of hybridizing to said polymorphic region;
- (e) DNA sequences which differ from any of the foregoing DNA sequences in codon sequence due to the degeneracy of the genetic code;
- (f) DNA sequences which are allelic variants of any of the foregoing DNA sequences; and
- (g) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

29. An isolated DNA sequence encoding a polymorphic region of an HLA-DR- $\beta$  chain locus of the human lymphocyte antigen complex, said DNA sequence being selected from the group consisting of:

- (a) DNA sequences encoding amino acids 8-14 of said locus;
- (b) DNA sequences encoding amino acids 26-32 of said locus;
- (c) DNA sequences encoding amino acids 72-78 of said locus;
- (d) DNA sequences which are portions of any one of the foregoing DNA sequences and which are capable of hybridizing to said polymorphic region;
- (e) DNA sequences which differ from any of the foregoing DNA sequences in codon sequence due to the degeneracy of the genetic code; and
- (f) DNA sequences which are fully complementary to any of the foregoing DNA sequences.

33. An isolated DNA sequence selected from the group consisting of:

- (a) DNA sequences encoding a majority of the region defined by amino acids:

- (i) 8-14,
- (ii) 26-32,
- (iii) 39-45, or
- (iv) 72-78

of the polypeptide coded for by DNA insert DR- $\beta$ -A, DR- $\beta$ -B or DR- $\beta$ -C;

(b) DNA sequences that are allelic variants of any of the foregoing DNA sequences; and

(c) DNA sequences that are complementary to any of the foregoing DNA sequences.